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(54) PROGRAMMABLE CONDITIONAL SIRNAS AND USES THEREOF

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(57)ABSTRACT

Disclosed herein are programmable, conditionally activated small interfering RNA constructs (Cond-siRNAs) and methods of making and using the same as therapeutic agents. The Cond-siRNA comprises a sensor strand, a core strand, and a guide strand, which crossover to form a sensor duplex and a RNAi duplex attached to each other to form a single structure. Upon binding an input strand to the sensor strand, the Cond-siRNA is activated and releases RNAi targeting a desired gene.

Specification includes a Sequence Listing.

First bridge Sensor **RNAi** ole strand Duplex Duplex Cond-siRNA docked to Giardia Dicer crystal structure Guide Sensor strand strand Second Sensor overhang (could be 3' or 5') bridge'